

**Joint Effects of Ambient Air Pollutants on Pediatric Asthma Emergency Department Visits in Atlanta,  
1998-2004**

Andrea Winquist, Ellen Kirrane, Mitch Klein, Matthew Strickland, Lyndsey A. Darrow, Stefanie Sarnat,  
Katherine Gass, James Mulholland, Armistead Russell, Paige Tolbert

**Supplemental Material**

## Model Details

The joint effect models for estimating the joint effects of  $p$  pollutants, without interactions, had the following form:

$$\begin{aligned}
 \text{Log}[E(Y)] = & \alpha + \sum_{i=1}^p \beta_i (\text{pollutant})_i & (1) \\
 & + \lambda[\log(\text{non} - \text{asthma pediatric resp ED visits})] + g(\gamma_1, \dots, \gamma_N; \text{day of season}) \\
 & + g(\delta_1, \dots, \delta_N; \text{dewpt}) + g(\eta_1, \dots, \eta_N; \text{min temp}) + \sum_j \xi_j (\text{year})_j \\
 & + \sum_k \nu_k (\text{month})_k + \sum_l \zeta_l (\text{day of week or holiday})_l + \sum_m \varphi_m (\text{hospital})_m \\
 & + \sum_n \tau_n (\text{max temp } ^\circ\text{C})_n + \sum_{jk} \psi_{jk} (\text{year})_j (\text{month})_k \\
 & + \sum_{kn} \omega_{kn} (\text{month})_k (\text{max temp } ^\circ\text{C})_n \\
 & + \sum_{kl} \vartheta_{kl} (\text{month})_k (\text{day of week or holiday})_l
 \end{aligned}$$

The dependent variable (Y) was the hospital-specific number of daily pediatric asthma ED visits. Analyses used the three-day moving average of pollutant concentrations (the average of the pollutant concentration on a given day (lag 0) and the previous two days (lags 1 and 2)) (*pollutant*), which was modeled as a linear term in our primary models. Models included a linear term for the logarithm of the daily number of non-asthma pediatric ED visits for acute upper respiratory infections (*non-asthma pediatric resp ED visits*); cubic polynomials ( $g$ ) for day-of-season, the moving average of dew point (lags 0-2) (*dewpt*), and the moving average of minimum temperature (lags 1 and 2) (*min temp*); indicator variables for year, month, day of week or holiday (with holidays having a separate indicator), hospital and same-day (lag 0) maximum temperature (for each degree Celsius) (*max temp °C*); and interaction terms between month and year, month and lag 0 maximum temperature, and month and day of week.

Joint effects were calculated for an interquartile-range (IQR) increment in each pollutant concentration, as the exponentiated sum (across the pollutants in the combination) of the product of each pollutant's model coefficient and that pollutant's IQR, using the following formulas:

$$\mathbf{L}'\boldsymbol{\beta} = \sum_{i=1}^p \beta_i (\text{pollutant IQR})_i \quad (2)$$

$$RR_{JE} = \exp[\mathbf{L}'\boldsymbol{\beta}] \quad (3)$$

where the joint effect of  $p$  pollutants ( $i=1, 2, \dots, p$ ) is represented by  $RR_{JE}$ ,  $\beta_i$  is the coefficient for a given pollutant ( $i$ ) from equation 1 above,  $\boldsymbol{\beta}$  is the vector of pollutant parameter estimates, and  $\mathbf{L}$  is the corresponding vector of pollutant IQRs. The 95% confidence interval for the joint effect was calculated (using an "estimate" statement in the SAS "genmod" procedure), according to the following formula:

$$se_{L'\beta} = \sqrt{\mathbf{L}'\hat{\boldsymbol{\Sigma}}\mathbf{L}} \quad (4)$$

$$95\% CI_{RR_{JE}} = (\exp(\mathbf{L}'\boldsymbol{\beta} - z_{1-\alpha/2}se_{L'\beta}), \exp(\mathbf{L}'\boldsymbol{\beta} + z_{1-\alpha/2}se_{L'\beta})) \quad (5)$$

Where  $\hat{\boldsymbol{\Sigma}}$  is the estimated covariance matrix of the estimates and  $\alpha = 0.05$ .

The joint effect models for estimating the joint effects of  $p$  pollutants with interactions had the following form:

$$\begin{aligned}
\text{Log}[E(Y)] = & \alpha + \sum_{i=1}^p \beta_i (\text{pollutant})_i + \sum_{i=2}^p \sum_{q=1}^{i-1} \beta_{iq} (\text{pollutant})_i * (\text{pollutant})_q & (6) \\
& + \lambda[\log(\text{non} - \text{asthma pediatric resp ED visits})] + g(\gamma_1, \dots, \gamma_N; \text{day of season}) \\
& + g(\delta_1, \dots, \delta_N; \text{dewpt}) + g(\eta_1, \dots, \eta_N; \text{min temp}) + \sum_j \xi_j (\text{year})_j \\
& + \sum_k \nu_k (\text{month})_k + \sum_l \zeta_l (\text{day of week or holiday})_l + \sum_m \varphi_m (\text{hospital})_m \\
& + \sum_n \tau_n (\text{max temp } ^\circ\text{C})_n + \sum_{jk} \psi_{jk} (\text{year})_j (\text{month})_k \\
& + \sum_{kn} \omega_{kn} (\text{month})_k (\text{max temp } ^\circ\text{C})_n \\
& + \sum_{kl} \vartheta_{kl} (\text{month})_k (\text{day of week or holiday})_l
\end{aligned}$$

Joint effects for the interaction models were calculated for an increment equal in magnitude to the interquartile-range (IQR) for each pollutant concentration, starting at each pollutant's 15<sup>th</sup>, 25<sup>th</sup>, or 35<sup>th</sup> percentile levels. For joint effects including interactions, the joint effect was calculated as the exponentiated sum (across the pollutants in the combination and their first-order interactions) of 1) the product of each pollutant's model coefficient and that pollutant's IQR, and 2) the product of the coefficient for each interaction term and the difference in the products of the specific pollutant levels being contrasted (represented by  $a$  and  $b$  in equation 7 below, where  $b=a+\text{IQR}$ ), using the following formulas:

$$\mathbf{L}'\boldsymbol{\beta} = \sum_{i=1}^p \beta_i (\text{pollutant IQR})_i + \sum_{i=2}^p \sum_{q=1}^{i-1} \beta_{iq} [(b_i * b_q) - (a_i * a_q)] \quad (7)$$

$$RR_{JE} = \exp[\mathbf{L}'\boldsymbol{\beta}] \quad (8)$$

where the joint effect of  $p$  pollutants ( $i=1, 2, \dots, p$ ) is represented by  $RR_{JE}$ ,  $\beta_i$  is the coefficient for a given pollutant ( $i$ ) from equation 6 above,  $\beta_{iq}$  is the coefficient for the interaction term for a given pollutant pair,  $\boldsymbol{\beta}$  is the vector of parameter estimates including estimates for pollutant  $s$  and interaction terms, and  $\mathbf{L}$  is the corresponding vector of pollutant IQRs (for pollutant betas) or differences in the product of contrasted pollutant levels (for interaction term betas). As in the no-interaction model, the 95% confidence interval for the joint effect was calculated (using an “estimate” statement in the SAS “genmod” procedure), according to the following formula:

$$se_{L'\boldsymbol{\beta}} = \sqrt{\mathbf{L}'\hat{\boldsymbol{\Sigma}}\mathbf{L}} \quad (9)$$

$$95\% CI_{RR_{JE}} = (\exp(\mathbf{L}'\boldsymbol{\beta} - z_{1-\alpha/2}se_{L'\boldsymbol{\beta}}), \exp(\mathbf{L}'\boldsymbol{\beta} + z_{1-\alpha/2}se_{L'\boldsymbol{\beta}})) \quad (10)$$

Where  $\hat{\boldsymbol{\Sigma}}$  is the estimated covariance matrix of the estimates and  $\alpha = 0.05$ .

Models used for estimating joint effects that included quadratic and cubic pollutant terms were similar to the model shown in equation 6 above except that quadratic and cubic terms for each pollutant were added to the model rather than interaction terms. The joint effect from these models was calculated as the exponentiated sum (across the pollutants in the combination and their quadratic and cubic terms) of 1) the product of each pollutant’s model coefficient and that pollutant’s IQR, and 2) the product of the coefficient for each quadratic and cubic term and the corresponding difference in the squared or cubed pollutant values for the specific pollutant levels being contrasted, in a manner similar to that shown in equation 7 above.

**eTable 1. Percentage of warm and cold season days with all pollutants in various quartiles of their respective year-round distributions,<sup>a</sup> Atlanta, 1998-2004**

Combination	Season	All Quartile 1	All Quartile 2	All Quartile 3	All Quartile 4	All not in same quartile
Oxidant Gases (O <sub>3</sub> , NO <sub>2</sub> , SO <sub>2</sub> )	WARM	3.6%	1.3%	1.3%	4.8%	89.0%
Secondary (O <sub>3</sub> , Secondary PM <sub>2.5</sub> <sup>b</sup> )	WARM	3.7%	5.4%	9.4%	26.4%	55.1%
Traffic (CO, NO <sub>2</sub> , EC)	WARM	11.5%	4.6%	4.2%	10.5%	69.2%
Power Plant (SO <sub>2</sub> , SO <sub>4</sub> <sup>2-</sup> )	WARM	4.9%	4.7%	7.0%	11.2%	72.2%
Criteria Pollutants (O <sub>3</sub> , CO, NO <sub>2</sub> , SO <sub>2</sub> , PM <sub>2.5</sub> )	WARM	1.8%	0.4%	0.5%	2.6%	94.7%
Oxidant Gases (O <sub>3</sub> , NO <sub>2</sub> , SO <sub>2</sub> )	COLD	4.5%	1.2%	2.0%	1.1%	91.2%
Secondary (O <sub>3</sub> , Secondary PM <sub>2.5</sub> <sup>b</sup> )	COLD	14.4%	9.0%	5.6%	1.5%	69.5%
Traffic (CO, NO <sub>2</sub> , EC)	COLD	9.1%	2.3%	3.0%	13.8%	71.8%
Power Plant (SO <sub>2</sub> , SO <sub>4</sub> <sup>2-</sup> )	COLD	8.4%	6.4%	5.4%	1.9%	77.9%
Criteria Pollutants (O <sub>3</sub> , CO, NO <sub>2</sub> , SO <sub>2</sub> , PM <sub>2.5</sub> )	COLD	1.7%	0.1%	0.2%	0.5%	97.5%

Definition of Abbreviations: O<sub>3</sub>=ozone, CO=carbon monoxide, NO<sub>2</sub>=nitrogen dioxide, SO<sub>2</sub>=sulfur dioxide, PM<sub>2.5</sub>=particulate matter less than 2.5 μm in diameter, EC=elemental carbon component of PM<sub>2.5</sub>, SO<sub>4</sub><sup>2-</sup>=sulfate component of PM<sub>2.5</sub>.

<sup>a</sup> Frequencies are calculated for percentage of warm or cold season days on which the various daily pollutant values were in various quartiles of the overall-year daily pollutant distribution.

<sup>b</sup> Secondary PM<sub>2.5</sub> was calculated as the sum of the concentrations of selected PM<sub>2.5</sub> components including sulfate, nitrate, and ammonium.

**eTable 2. Primary joint effect model effect estimates, Atlanta, 1998-2004, calculated for an inter-quartile range<sup>a</sup> change in the 3-day moving average of each pollutant**

Effect	Season	Model Specifications	Joint Effect Estimate		Interaction p-value <sup>b</sup>
			RR (95% CI)	p-value	
Oxidant Gases (O <sub>3</sub> , NO <sub>2</sub> , SO <sub>2</sub> )	WARM	Model including only pollutants in joint effect, no interactions	1.0981 (1.0440-1.1550)	0.0003	NA
		Model including only pollutants in joint effect, with first order interactions	1.0997 (1.0432-1.1593)	0.0004	0.9748
Secondary (O <sub>3</sub> , Secondary PM <sub>2.5</sub> <sup>c</sup> )	WARM	Model including only pollutants in joint effect, no interactions	1.0851 (1.0354-1.1372)	0.0006	NA
		Model including only pollutants in joint effect, with first order interactions	1.0853 (1.0281-1.1457)	0.0031	0.9900
Traffic (CO, NO <sub>2</sub> , EC)	WARM	Model including only pollutants in joint effect, no interactions	1.1108 (1.0621-1.1616)	<0.0001	NA
		Model including only pollutants in joint effect, with first order interactions	1.1332 (1.0768-1.1924)	<0.0001	0.4209
Power Plant (SO <sub>2</sub> , SO <sub>4</sub> <sup>2-</sup> )	WARM	Model including only pollutants in joint effect, no interactions	1.0574 (1.0211-1.0950)	0.0018	NA
		Model including only pollutants in joint effect, with first order interactions	1.0829 (1.0374-1.1304)	0.0003	0.0598
Criteria Pollutants (O <sub>3</sub> , CO, NO <sub>2</sub> , SO <sub>2</sub> , PM <sub>2.5</sub> )	WARM	Model including only pollutants in joint effect, no interactions	1.1289 (1.0573-1.2053)	0.0003	NA
		Model including only pollutants in joint effect, with first order interactions	1.1540 (1.0698-1.2449)	0.0002	0.3314
Oxidant Gases (O <sub>3</sub> , NO <sub>2</sub> , SO <sub>2</sub> )	COLD	Model including only pollutants in joint effect, no interactions	1.0539 (0.9851-1.1275)	0.1274	NA
		Model including only pollutants in joint effect, with first order interactions	1.0307 (0.9548-1.1126)	0.4386	0.0088
Secondary (O <sub>3</sub> , Secondary PM <sub>2.5</sub> <sup>c</sup> )	COLD	Model including only pollutants in joint effect, no interactions	1.0617 (0.9918-1.1366)	0.0850	NA
		Model including only pollutants in joint effect, with first order interactions	1.0457 (0.9745-1.1222)	0.2138	0.0890
Traffic (CO, NO <sub>2</sub> , EC)	COLD	Model including only pollutants in joint effect, no interactions	1.0194 (0.9896-1.0500)	0.2045	NA
		Model including only pollutants in joint effect, with first order interactions	1.0182 (0.9799-1.0579)	0.3565	0.9941
Power Plant (SO <sub>2</sub> , SO <sub>4</sub> <sup>2-</sup> )	COLD	Model including only pollutants in joint effect, no interactions	0.9810 (0.9398-1.0240)	0.3803	NA
		Model including only pollutants in joint effect, with first order interactions	0.9845 (0.9428-1.0280)	0.4788	0.2658
Criteria Pollutants (O <sub>3</sub> , CO, NO <sub>2</sub> , SO <sub>2</sub> , PM <sub>2.5</sub> )	COLD	Model including only pollutants in joint effect, no interactions	1.0397 (0.9670-1.1178)	0.2926	NA
		Model including only pollutants in joint effect, with first order interactions	0.9943 (0.9026-1.0953)	0.9077	0.0002

Definition of Abbreviations: O<sub>3</sub>=ozone, CO=carbon monoxide, NO<sub>2</sub>=nitrogen dioxide, SO<sub>2</sub>=sulfur dioxide, PM<sub>2.5</sub>=particulate matter less than 2.5 µm in diameter, EC=elemental carbon component of PM<sub>2.5</sub>, SO<sub>4</sub><sup>2-</sup>=sulfate component of PM<sub>2.5</sub>, RR=Rate Ratio

<sup>a</sup> For comparability with previous analyses,<sup>1</sup> the analysis used the year-round IQR during 1993-2004 for O<sub>3</sub>, NO<sub>2</sub>, CO and SO<sub>2</sub>, and during August 1, 1998-December 31, 2004 for PM<sub>2.5</sub> and PM<sub>2.5</sub> components. The IQRs used in the analysis were: O<sub>3</sub> 29.18 ppb, CO 0.66 ppm, NO<sub>2</sub> 12.87 ppb, SO<sub>2</sub> 10.51 ppb, PM<sub>2.5</sub> 9.18 µg/m<sup>3</sup>, EC 0.69 µg/m<sup>3</sup>, SO<sub>4</sub><sup>2-</sup> 3.45 µg/m<sup>3</sup>, Secondary PM<sub>2.5</sub> 4.52 µg/m<sup>3</sup>, NO<sub>x</sub> 42.70 ppb.

<sup>b</sup> Interaction test was a likelihood ratio test, testing the hypothesis that the coefficients for all interaction terms equal 0, performed using a contrast statement in SAS PROC GENMOD.

<sup>c</sup> Secondary PM<sub>2.5</sub> was calculated as the sum of the concentrations of selected PM<sub>2.5</sub> components including sulfate, nitrate, and ammonium.

**eTable 3. Single-pollutant model effect estimates, Atlanta, warm and cold seasons , 1998-2004, calculated for an inter-quartile range<sup>a</sup> change in the 3-day moving average of in each pollutant**

Effect	Season	Model Specifications	Effect Estimate	
			RR (95% CI)	p-value
O <sub>3</sub>	WARM	Single Pollutant Model	1.0778 (1.0308-1.1270)	0.0010
CO	WARM	Single Pollutant Model	1.1076 (1.0597-1.1576)	0.0000
NO <sub>2</sub>	WARM	Single Pollutant Model	1.0868 (1.0536-1.1211)	0.0000
SO <sub>2</sub>	WARM	Single Pollutant Model	1.0426 (1.0112-1.0750)	0.0074
PM <sub>2.5</sub>	WARM	Single Pollutant Model	1.0430 (1.0165-1.0703)	0.0014
EC	WARM	Single Pollutant Model	1.0401 (1.0097-1.0714)	0.0095
SO <sub>4</sub> <sup>2-</sup>	WARM	Single Pollutant Model	1.0261 (1.0041-1.0487)	0.0200
Secondary PM <sub>2.5</sub> <sup>b</sup>	WARM	Single Pollutant Model	1.0263 (1.0049-1.0481)	0.0159
O <sub>3</sub>	COLD	Single Pollutant Model	1.0702 (1.0079-1.1364)	0.0266
CO	COLD	Single Pollutant Model	1.0150 (0.9869-1.0439)	0.2993
NO <sub>2</sub>	COLD	Single Pollutant Model	1.0193 (0.9917-1.0478)	0.1723
SO <sub>2</sub>	COLD	Single Pollutant Model	0.9841 (0.9578-1.0111)	0.2446
PM <sub>2.5</sub>	COLD	Single Pollutant Model	1.0045 (0.9785-1.0311)	0.7372
EC	COLD	Single Pollutant Model	1.0032 (0.9814-1.0254)	0.7787
SO <sub>4</sub> <sup>2-</sup>	COLD	Single Pollutant Model	0.9907 (0.9538-1.0290)	0.6306
Secondary PM <sub>2.5</sub> <sup>b</sup>	COLD	Single Pollutant Model	0.9840 (0.9530-1.0160)	0.3221

Definition of Abbreviations: O<sub>3</sub>=ozone, CO=carbon monoxide, NO<sub>2</sub>=nitrogen dioxide, SO<sub>2</sub>=sulfur dioxide, PM<sub>2.5</sub>=particulate matter less than 2.5 μm in diameter, EC=elemental carbon component of PM<sub>2.5</sub>, SO<sub>4</sub><sup>2-</sup>=sulfate component of PM<sub>2.5</sub>, RR=Rate Ratio

<sup>a</sup> For comparability with previous analyses,<sup>1</sup> the analysis used the year-round IQR during 1993-2004 for O<sub>3</sub>, NO<sub>2</sub>, CO and SO<sub>2</sub>, and during August 1, 1998-December 31, 2004 for PM<sub>2.5</sub> and PM<sub>2.5</sub> components. The IQRs used in the analysis were: O<sub>3</sub> 29.18 ppb, CO 0.66 ppm, NO<sub>2</sub> 12.87 ppb, SO<sub>2</sub> 10.51 ppb, PM<sub>2.5</sub> 9.18 μg/m<sup>3</sup>, EC 0.69 μg/m<sup>3</sup>, SO<sub>4</sub><sup>2-</sup> 3.45 μg/m<sup>3</sup>, Secondary PM<sub>2.5</sub> 4.52 μg/m<sup>3</sup>.

<sup>b</sup> Secondary PM<sub>2.5</sub> was calculated as the sum of the concentrations of selected PM<sub>2.5</sub> components including sulfate, nitrate, and ammonium.



**Table 4. Pollutant-specific parameter and concurrency estimates, primary models, Atlanta, 1998-2004, Warm Season**

Effect	Model Specifications	Pollutant*	Unit	Coefficient per 1 unit change	p-value	Concurrency: all variables	Concurrency: pollutants only	
O <sub>3</sub>	Single Pollutant Model	O <sub>3</sub>	ppb	0.0026	0.0010	0.874	NA	
CO	Single Pollutant Model	CO	ppm	0.1548	0.0000	0.841	NA	
NO <sub>2</sub>	Single Pollutant Model	NO <sub>2</sub>	ppb	0.0065	0.0000	0.812	NA	
SO <sub>2</sub>	Single Pollutant Model	SO <sub>2</sub>	ppb	0.0040	0.0074	0.609	NA	
PM <sub>2.5</sub>	Single Pollutant Model	PM <sub>2.5</sub>	µg/m <sup>3</sup>	0.0046	0.0014	0.734	NA	
EC	Single Pollutant Model	EC	µg/m <sup>3</sup>	0.0569	0.0095	0.825	NA	
SO <sub>4</sub> <sup>2-</sup>	Single Pollutant Model	SO <sub>4</sub> <sup>2-</sup>	µg/m <sup>3</sup>	0.0075	0.0200	0.726	NA	
Secondary PM <sub>2.5</sub> <sup>a</sup>	Single Pollutant Model	Secondary PM <sub>2.5</sub> <sup>a</sup>	µg/m <sup>3</sup>	0.0057	0.0159	0.727	NA	
Oxidant Gases	Model including only pollutants in joint effect, no interactions	O <sub>3</sub>	ppb	0.0003	0.7949	0.923	0.566	
		NO <sub>2</sub>	ppb	0.0059	0.0004	0.901	0.646	
		SO <sub>2</sub>	ppb	0.0010	0.5495	0.686	0.443	
	Model including only pollutants in joint effect, with first order interactions	O <sub>3</sub>	ppb	0.0006	0.7216	0.972	0.944	
		NO <sub>2</sub>	ppb	0.0065	0.1057	0.985	0.972	
		SO <sub>2</sub>	ppb	-0.0010	0.8756	0.982	0.976	
		O <sub>3</sub> *NO <sub>2</sub>	ppb <sup>2</sup>	0.0000	0.7608	0.992	0.986	
Secondary	Model including only pollutants in joint effect, no interactions	O <sub>3</sub>	ppb	0.0026	0.0091	0.927	0.586	
		Secondary PM <sub>2.5</sub> <sup>a</sup>	µg/m <sup>3</sup>	0.0014	0.6247	0.843	0.583	
	Model including only pollutants in joint effect, with first order interactions	O <sub>3</sub>	ppb	0.0026	0.0621	0.960	0.897	
		Secondary PM <sub>2.5</sub> <sup>a</sup>	µg/m <sup>3</sup>	0.0015	0.8375	0.977	0.960	
		O <sub>3</sub> *Secondary PM <sub>2.5</sub> <sup>a</sup>	ppb*	0.0000	0.9900	0.987	0.978	
	Traffic	Model including only pollutants in joint effect, no interactions	CO	ppm	0.1151	0.0377	0.940	0.866
			NO <sub>2</sub>	ppb	0.0052	0.0044	0.916	0.777
EC			µg/m <sup>3</sup>	-0.0549	0.0898	0.918	0.839	
Model including only pollutants in joint effect, with first order interactions		CO	ppm	0.1257	0.3404	0.990	0.984	
		NO <sub>2</sub>	ppb	0.0047	0.2399	0.984	0.973	
		EC	µg/m <sup>3</sup>	0.0380	0.6552	0.989	0.981	
		CO*NO <sub>2</sub>	ppm*ppb	0.0017	0.6877	0.996	0.994	
Power Plant	Model including only pollutants in joint effect, no interactions	SO <sub>2</sub>	ppb	0.0033	0.0320	0.634	0.256	
		SO <sub>4</sub> <sup>2-</sup>	µg/m <sup>3</sup>	0.0060	0.0667	0.738	0.261	
	Model including only pollutants in joint effect, with first order interactions	SO <sub>2</sub>	ppb	0.0099	0.0096	0.950	0.927	
		SO <sub>4</sub> <sup>2-</sup>	µg/m <sup>3</sup>	0.0150	0.0096	0.927	0.890	
		SO <sub>2</sub> *SO <sub>4</sub> <sup>2-</sup>	ppb*	-0.0010	0.0602	0.971	0.961	
	Criteria Pollutants	Model including only pollutants in joint effect, no interactions	O <sub>3</sub>	ppb	0.0008	0.5182	0.953	0.792
			CO	ppm	0.0854	0.0909	0.929	0.820
NO <sub>2</sub>			ppb	0.0032	0.1460	0.942	0.864	
SO <sub>2</sub>			ppb	0.0007	0.6979	0.693	0.467	
PM <sub>2.5</sub>			µg/m <sup>3</sup>	-0.0006	0.7814	0.900	0.767	
Model including only pollutants in joint effect, with first order interactions		O <sub>3</sub>	ppb	-0.0034	0.1857	0.989	0.979	
		CO	ppm	0.0877	0.5605	0.992	0.988	
		NO <sub>2</sub>	ppb	0.0099	0.1427	0.995	0.991	
		SO <sub>2</sub>	ppb	-0.0039	0.5749	0.984	0.978	
		PM <sub>2.5</sub>	µg/m <sup>3</sup>	0.0012	0.8477	0.990	0.985	
		O <sub>3</sub> *CO	ppb*ppm	0.0016	0.5689	0.996	0.994	
		O <sub>3</sub> *NO <sub>2</sub>	ppb <sup>2</sup>	0.0000	0.7481	0.997	0.996	
		O <sub>3</sub> *SO <sub>2</sub>	ppb <sup>2</sup>	0.0002	0.2309	0.993	0.991	
		O <sub>3</sub> *PM <sub>2.5</sub>	ppb*	0.0001	0.2531	0.996	0.993	
		CO*NO <sub>2</sub>	ppm*ppb	-0.0061	0.1016	0.994	0.991	
CO*SO <sub>2</sub>	ppm*ppb	0.0076	0.2357	0.993	0.991			
CO*PM <sub>2.5</sub>	ppm*	0.0015	0.8420	0.997	0.996			
NO <sub>2</sub> *SO <sub>2</sub>	ppb <sup>2</sup>	0.0002	0.5174	0.995	0.993			
NO <sub>2</sub> *PM <sub>2.5</sub>	ppb*	-0.0001	0.7281	0.998	0.997			
SO <sub>2</sub> *PM <sub>2.5</sub>	ppb*	-0.0008	0.0361	0.994	0.992			

Abbreviations: O<sub>3</sub>=ozone, CO=carbon monoxide, NO<sub>2</sub>=nitrogen dioxide, SO<sub>2</sub>=sulfur dioxide, PM<sub>2.5</sub>=particulate matter < 2.5 µm in diameter, EC=elemental carbon component of PM<sub>2.5</sub>, SO<sub>4</sub><sup>2-</sup>=sulfate component of PM<sub>2.5</sub>, ppb=parts per billion, ppm=parts per million, µg/m<sup>3</sup>=micrograms per cubic millimeter.

<sup>a</sup>Secondary PM<sub>2.5</sub> was calculated as the sum of the concentrations of the PM<sub>2.5</sub> components sulfate, nitrate, and ammonium.

**Table 5. Pollutant-specific parameter and concurrency estimates, primary models, Atlanta, 1998-2004, Cold Season**

Effect	Model Specifications	Pollutant*	Unit	Coefficient per 1 unit change	p-value	Concurrency: all variables	Concurrency: pollutants only	
O <sub>3</sub>	Single Pollutant Model	O <sub>3</sub>	ppb	0.0023	0.0266	0.905	NA	
CO	Single Pollutant Model	CO	ppm	0.0225	0.2993	0.740	NA	
NO <sub>2</sub>	Single Pollutant Model	NO <sub>2</sub>	ppb	0.0015	0.1723	0.740	NA	
SO <sub>2</sub>	Single Pollutant Model	SO <sub>2</sub>	ppb	-0.0015	0.2446	0.721	NA	
PM <sub>2.5</sub>	Single Pollutant Model	PM <sub>2.5</sub>	µg/m <sup>3</sup>	0.0005	0.7372	0.711	NA	
EC	Single Pollutant Model	EC	µg/m <sup>3</sup>	0.0046	0.7787	0.776	NA	
SO <sub>4</sub> <sup>2-</sup>	Single Pollutant Model	SO <sub>4</sub> <sup>2-</sup>	µg/m <sup>3</sup>	-0.0027	0.6306	0.748	NA	
Secondary PM <sub>2.5</sub> <sup>a</sup>	Single Pollutant Model	Secondary PM <sub>2.5</sub> <sup>a</sup>	µg/m <sup>3</sup>	-0.0036	0.3221	0.700	NA	
Oxidant Gases	Model including only pollutants in joint effect, no interactions	O <sub>3</sub>	ppb	0.0019	0.0935	0.917	0.398	
		NO <sub>2</sub>	ppb	0.0012	0.3085	0.790	0.531	
		SO <sub>2</sub>	ppb	-0.0017	0.2175	0.751	0.422	
	Model including only pollutants in joint effect, with first order interactions	O <sub>3</sub>	ppb	0.0011	0.6493	0.982	0.965	
		NO <sub>2</sub>	ppb	0.0006	0.8541	0.976	0.964	
		SO <sub>2</sub>	ppb	0.0127	0.0059	0.981	0.972	
		O <sub>3</sub> *NO <sub>2</sub>	ppb <sup>2</sup>	0.0001	0.1583	0.990	0.985	
Secondary	Model including only pollutants in joint effect, no interactions	O <sub>3</sub>	ppb	0.0026	0.0133	0.908	0.166	
		Secondary PM <sub>2.5</sub> <sup>a</sup>	µg/m <sup>3</sup>	-0.0038	0.2877	0.700	0.170	
	Model including only pollutants in joint effect, with first order interactions	O <sub>3</sub>	ppb	0.0056	0.0059	0.975	0.940	
		Secondary PM <sub>2.5</sub> <sup>a</sup>	µg/m <sup>3</sup>	0.0134	0.2135	0.971	0.949	
		O <sub>3</sub> *Secondary PM <sub>2.5</sub> <sup>a</sup>	ppb*	-0.0005	0.0894	0.985	0.975	
	Traffic	Model including only pollutants in joint effect, no interactions	CO	ppm	-0.0074	0.8466	0.920	0.828
			NO <sub>2</sub>	ppb	0.0028	0.1410	0.922	0.739
EC			µg/m <sup>3</sup>	-0.0180	0.4414	0.897	0.765	
Model including only pollutants in joint effect, with first order interactions		CO	ppm	0.0154	0.8940	0.992	0.986	
		NO <sub>2</sub>	ppb	0.0025	0.5478	0.984	0.964	
		EC	µg/m <sup>3</sup>	-0.0356	0.6753	0.992	0.982	
		CO*NO <sub>2</sub>	ppm*ppb	-0.0004	0.9117	0.996	0.993	
Power Plant	Model including only pollutants in joint effect, no interactions	SO <sub>2</sub>	ppb	-0.0013	0.3334	0.746	0.053	
		SO <sub>4</sub> <sup>2-</sup>	µg/m <sup>3</sup>	-0.0015	0.7984	0.762	0.069	
	Model including only pollutants in joint effect, with first order interactions	SO <sub>2</sub>	ppb	-0.0042	0.1500	0.950	0.915	
		SO <sub>4</sub> <sup>2-</sup>	µg/m <sup>3</sup>	-0.0113	0.2841	0.933	0.887	
		SO <sub>2</sub> *SO <sub>4</sub> <sup>2-</sup>	ppb*	0.0009	0.2654	0.963	0.947	
	Criteria Pollutants	Model including only pollutants in joint effect, no interactions	O <sub>3</sub>	ppb	0.0017	0.1539	0.922	0.456
			CO	ppm	0.0078	0.8306	0.913	0.783
NO <sub>2</sub>			ppb	0.0022	0.2942	0.933	0.815	
SO <sub>2</sub>			ppb	-0.0020	0.1536	0.755	0.446	
PM <sub>2.5</sub>			µg/m <sup>3</sup>	-0.0024	0.2096	0.838	0.642	
Model including only pollutants in joint effect, with first order interactions		O <sub>3</sub>	ppb	-0.0007	0.8073	0.988	0.977	
		CO	ppm	0.2354	0.0833	0.994	0.989	
		NO <sub>2</sub>	ppb	-0.0048	0.4776	0.994	0.990	
		SO <sub>2</sub>	ppb	0.0093	0.0818	0.985	0.979	
		PM <sub>2.5</sub>	µg/m <sup>3</sup>	-0.0076	0.2588	0.987	0.981	
		O <sub>3</sub> *CO	ppb*ppm	-0.0135	0.0000	0.995	0.991	
		O <sub>3</sub> *NO <sub>2</sub>	ppb <sup>2</sup>	0.0004	0.0008	0.996	0.993	
		O <sub>3</sub> *SO <sub>2</sub>	ppb <sup>2</sup>	-0.0002	0.0425	0.981	0.972	
		O <sub>3</sub> *PM <sub>2.5</sub>	ppb*	0.0003	0.0814	0.995	0.992	
Secondary PM <sub>2.5</sub> <sup>a</sup>	CO*NO <sub>2</sub>	ppm*ppb	0.0089	0.0128	0.996	0.994		
	CO*SO <sub>2</sub>	ppm*ppb	-0.0081	0.1094	0.990	0.986		
	CO*PM <sub>2.5</sub>	ppm*	0.0043	0.5036	0.997	0.994		
	NO <sub>2</sub> *SO <sub>2</sub>	ppb <sup>2</sup>	-0.0004	0.1216	0.994	0.992		
	NO <sub>2</sub> *PM <sub>2.5</sub>	ppb*	-0.0008	0.0199	0.998	0.996		
SO <sub>2</sub> *PM <sub>2.5</sub>	ppb*	0.0011	0.0075	0.992	0.989			

Abbreviations: O<sub>3</sub>=ozone, CO=carbon monoxide, NO<sub>2</sub>=nitrogen dioxide, SO<sub>2</sub>=sulfur dioxide, PM<sub>2.5</sub>=particulate matter < 2.5 µm in diameter, EC=elemental carbon component of PM<sub>2.5</sub>, SO<sub>4</sub><sup>2-</sup>=sulfate component of PM<sub>2.5</sub>, ppb=parts per billion, ppm=parts per million, µg/m<sup>3</sup>=micrograms per cubic millimeter.

<sup>a</sup>Secondary PM<sub>2.5</sub> was calculated as the sum of the concentrations of the PM<sub>2.5</sub> components sulfate, nitrate, and ammonium.

eTable 6. Variance-Covariance matrices for primary joint effect models, Atlanta, 1998-2004, Warm Season

		O <sub>3</sub>	CO	NO <sub>2</sub>	SO <sub>2</sub>	PM <sub>2.5</sub>	EC	SO <sub>4</sub> <sup>2-</sup>	Secondary PM <sub>2.5</sub> <sup>a</sup>
Oxidant Gases	O <sub>3</sub>	9.5546E-07		-9.6111E-07	1.6987E-07				
	NO <sub>2</sub>	-9.6111E-07		2.7883E-06	-1.0764E-06				
	SO <sub>2</sub>	1.6987E-07		-1.0764E-06	2.6953E-06				
Secondary	O <sub>3</sub>	9.7868E-07							-1.6393E-06
	Secondary PM <sub>2.5</sub> <sup>a</sup>	-1.6393E-06							8.4099E-06
Traffic	CO		3.0670E-03	-4.7960E-05			-8.6823E-04		
	NO <sub>2</sub>		-4.7960E-05	3.3395E-06			-1.4614E-05		
	EC		-8.6823E-04	-1.4614E-05			1.0455E-03		
Power Plant	SO <sub>2</sub>				2.4133E-06			-1.0574E-06	
	SO <sub>4</sub> <sup>2-</sup>				-1.0574E-06			1.0806E-05	
Criteria Pollutants	O <sub>3</sub>	1.4406E-06	1.6754E-05	-1.1306E-06	1.9821E-07	-1.4534E-06			
	CO	1.6754E-05	2.5525E-03	-6.5342E-05	-3.0834E-06	-2.9971E-05			
	NO <sub>2</sub>	-1.1306E-06	-6.5342E-05	4.7764E-06	-1.0238E-06	-1.9850E-07			
	SO <sub>2</sub>	1.9821E-07	-3.0834E-06	-1.0238E-06	2.8149E-06	-9.1418E-08			
	PM <sub>2.5</sub>	-1.4534E-06	-2.9971E-05	-1.9850E-07	-9.1418E-08	4.6962E-06			

Definition of Abbreviations: O<sub>3</sub>=ozone, CO=carbon monoxide, NO<sub>2</sub>=nitrogen dioxide, SO<sub>2</sub>=sulfur dioxide, PM<sub>2.5</sub>=particulate matter less than 2.5 μm in diameter, EC=elemental carbon component of PM<sub>2.5</sub>, SO<sub>4</sub><sup>2-</sup>=sulfate component of PM<sub>2.5</sub>.

<sup>a</sup>Secondary PM<sub>2.5</sub> was calculated as the sum of the concentrations of selected PM<sub>2.5</sub> components including nitrate (NO<sub>3</sub><sup>-</sup>), ammonium (NH<sub>4</sub><sup>+</sup>), and sulfate (SO<sub>4</sub><sup>2-</sup>).

**eTable 7. Variance-Covariance matrices for primary joint effect models, Atlanta, 1998-2004, Cold Season**

		O <sub>3</sub>	CO	NO <sub>2</sub>	SO <sub>2</sub>	PM <sub>2.5</sub>	EC	SO <sub>4</sub> <sup>2-</sup>	Secondary PM <sub>2.5</sub> <sup>a</sup>
<b>Oxidant Gases</b>	<b>O<sub>3</sub></b>	1.2486E-06		-4.5664E-07	2.4593E-07				
	<b>NO<sub>2</sub></b>	-4.5664E-07		1.4460E-06	-4.9985E-07				
	<b>SO<sub>2</sub></b>	2.4593E-07		-4.9985E-07	1.9131E-06				
<b>Secondary</b>	<b>O<sub>3</sub></b>	1.1417E-06							-1.1427E-07
	<b>Secondary PM<sub>2.5</sub><sup>a</sup></b>	-1.1427E-07							1.3045E-05
<b>Traffic</b>	<b>CO</b>		1.4512E-03	-4.5887E-05			-2.7387E-04		
	<b>NO<sub>2</sub></b>		-4.5887E-05	3.7117E-06			-1.4109E-05		
	<b>EC</b>		-2.7387E-04	-1.4109E-05			5.4888E-04		
<b>Power Plant</b>	<b>SO<sub>2</sub></b>				1.9323E-06			-1.7643E-06	
	<b>SO<sub>4</sub><sup>2-</sup></b>				-1.7643E-06			3.3063E-05	
<b>Criteria Pollutants</b>	<b>O<sub>3</sub></b>	1.3608E-06	5.3877E-06	-7.1054E-07	2.3490E-07	1.2589E-08			
	<b>CO</b>	5.3877E-06	1.3210E-03	-5.3425E-05	-1.1995E-06	-2.2904E-06			
	<b>NO<sub>2</sub></b>	-7.1054E-07	-5.3425E-05	4.4393E-06	-5.2562E-07	-1.5992E-06			
	<b>SO<sub>2</sub></b>	2.3490E-07	-1.1995E-06	-5.2562E-07	2.0331E-06	9.1886E-08			
	<b>PM<sub>2.5</sub></b>	1.2589E-08	-2.2904E-06	-1.5992E-06	9.1886E-08	3.5909E-06			

Definition of Abbreviations: O<sub>3</sub>=ozone, CO=carbon monoxide, NO<sub>2</sub>=nitrogen dioxide, SO<sub>2</sub>=sulfur dioxide, PM<sub>2.5</sub>=particulate matter less than 2.5 μm in diameter, EC=elemental carbon component of PM<sub>2.5</sub>, SO<sub>4</sub><sup>2-</sup>=sulfate component of PM<sub>2.5</sub>.

<sup>a</sup>Secondary PM<sub>2.5</sub> was calculated as the sum of the concentrations of selected PM<sub>2.5</sub> components including nitrate (NO<sub>3</sub><sup>-</sup>), ammonium (NH<sub>4</sub><sup>+</sup>), and sulfate (SO<sub>4</sub><sup>2-</sup>).

**eTable 8. Sensitivity analyses for joint effect calculations, Atlanta, 1998-2004, calculated for an inter-quartile range<sup>a</sup> change in the 3-day moving average of in each pollutant**

Effect	Season	Model Specifications	Joint Effect Estimate <sup>b</sup>		Interaction or Non-linear term p-value <sup>c</sup>
			RR (95% CI)	p-value	
Oxidant Gases (O <sub>3</sub> , NO <sub>2</sub> , SO <sub>2</sub> )	WARM	Model including pollutants in joint effect, also controlling for CO and PM <sub>2.5</sub> , no interactions	1.0729 (0.9990-1.1522)	0.0531	NA
		Model including only pollutants in joint effect, with linear, quadratic and cubic terms for each pollutant	1.0957 (1.0297-1.1658)	0.0039	0.8487
Secondary (O <sub>3</sub> , Secondary PM <sub>2.5</sub> <sup>d</sup> )	WARM	Model including pollutants in joint effect, also controlling for CO, NO <sub>2</sub> , and SO <sub>2</sub> , no interactions	1.0206 (0.9628-1.0820)	0.4929	NA
		Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	1.0855 (1.0233-1.1514)	0.0064	0.7532
Traffic (CO, NO <sub>2</sub> , EC)	WARM	Model including pollutants in joint effect, also controlling for O <sub>3</sub> , SO <sub>2</sub> and SO <sub>4</sub> <sup>2-</sup> , no interactions	1.1113 (1.0523-1.1735)	0.0001	NA
		Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	1.0900 (1.0279-1.1559)	0.0040	0.0262
Power Plant (SO <sub>2</sub> , SO <sub>4</sub> <sup>2-</sup> )	WARM	Model including pollutants in joint effect, also controlling for O <sub>3</sub> , CO, NO <sub>2</sub> and EC, no interactions	1.0033 (0.9613-1.0472)	0.8802	NA
		Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	1.0634 (1.0060-1.1240)	0.0299	0.7920
Criteria Pollutants (O <sub>3</sub> , CO, NO <sub>2</sub> , SO <sub>2</sub> , PM <sub>2.5</sub> )	WARM	Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	1.1149 (1.0322-1.2041)	0.0056	0.8846
Oxidant Gases (O <sub>3</sub> , NO <sub>2</sub> , SO <sub>2</sub> )	COLD	Model including pollutants in joint effect, also controlling for CO and PM <sub>2.5</sub> , no interactions	1.0572 (0.9778-1.143)	0.1628	NA
		Model including only pollutants in joint effect, with linear, quadratic and cubic terms for each pollutant	1.1323 (1.0324-1.2418)	0.0084	0.1948
Secondary (O <sub>3</sub> , Secondary PM <sub>2.5</sub> <sup>d</sup> )	COLD	Model including pollutants in joint effect, also controlling for CO, NO <sub>2</sub> , and SO <sub>2</sub> , no interactions	1.0333 (0.9574-1.1153)	0.3999	NA
		Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	1.1852 (1.0877-1.2914)	0.0001	0.0004
Traffic (CO, NO <sub>2</sub> , EC)	COLD	Model including pollutants in joint effect, also controlling for O <sub>3</sub> , SO <sub>2</sub> and SO <sub>4</sub> <sup>2-</sup> , no interactions	1.0242 (0.9912-1.0584)	0.1525	NA
		Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	1.0076 (0.9665-1.0505)	0.7216	0.9279
Power Plant (SO <sub>2</sub> , SO <sub>4</sub> <sup>2-</sup> )	COLD	Model including pollutants in joint effect, also controlling for O <sub>3</sub> , CO, NO <sub>2</sub> and EC, no interactions	0.9583 (0.9145-1.0042)	0.0743	NA
		Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	1.0258 (0.9727-1.0818)	0.3468	0.0118
Criteria Pollutants (O <sub>3</sub> , CO, NO <sub>2</sub> , SO <sub>2</sub> , PM <sub>2.5</sub> )	COLD	Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	1.1322 (1.0252-1.2503)	0.0142	0.0295

Abbreviations: O<sub>3</sub>=ozone, CO=carbon monoxide, NO<sub>2</sub>=nitrogen dioxide, SO<sub>2</sub>=sulfur dioxide, PM<sub>2.5</sub>=particulate matter < 2.5 µm in diameter, EC=elemental carbon component of PM<sub>2.5</sub>, SO<sub>4</sub><sup>2-</sup>=sulfate component of PM<sub>2.5</sub>, RR=Rate Ratio

<sup>a</sup> For comparability with previous analyses,<sup>1</sup> the analysis used the year-round IQR during 1993-2004 for O<sub>3</sub>, NO<sub>2</sub>, CO and SO<sub>2</sub>, and during August 1, 1998-December 31, 2004 for PM<sub>2.5</sub> and PM<sub>2.5</sub> components. The IQRs used in the analysis were: O<sub>3</sub> 29.18 ppb, CO 0.66 ppm, NO<sub>2</sub> 12.87 ppb, SO<sub>2</sub> 10.51 ppb, PM<sub>2.5</sub> 9.18 µg/m<sup>3</sup>, EC 0.69 µg/m<sup>3</sup>, SO<sub>4</sub><sup>2-</sup> 3.45 µg/m<sup>3</sup>, Secondary PM<sub>2.5</sub> 4.52 µg/m<sup>3</sup>, NO<sub>x</sub> 42.7 ppb, RR=Rate Ratio

<sup>b</sup> In interaction models, the joint effects were calculated for an IQR change from the 25<sup>th</sup> percentile level to the 75<sup>th</sup> percentile level for each pollutant.

<sup>c</sup> The test for the interaction terms and non-linear terms (quadratic and cubic terms) was a likelihood ratio test, testing the hypothesis that the coefficients for all interaction terms, or all quadratic and cubic terms, equal 0, performed using a contrast statement in SAS PROC GENMOD.

<sup>d</sup> Secondary PM<sub>2.5</sub> was calculated as the sum of the concentrations of the PM<sub>2.5</sub> components sulfate, nitrate, and ammonium.

**eTable 9. Pollutant-specific parameter estimates from sensitivity analyses, Atlanta, 1998-2004, Warm Season**

Joint Effect	Model Specifications	Pollutant <sup>a</sup>	Unit	Coefficient per 1 unit change	p-value
Oxidant Gases	Model including pollutants in joint effect, also controlling for CO and PM <sub>2.5</sub> , no interactions	O <sub>3</sub>	ppb	0.0008	0.5182
		CO	ppm	0.0854	0.0909
		NO <sub>2</sub>	ppb	0.0032	0.1460
		SO <sub>2</sub>	ppb	0.0143	0.6979
		PM <sub>2.5</sub>	µg/m <sup>3</sup>	-0.0006	0.7814
	Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	O <sub>3</sub>	ppb	-0.0096	0.1990
		(O <sub>3</sub> ) <sup>2</sup>	ppb <sup>2</sup>	0.0002	0.2043
		(O <sub>3</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.2376
		NO <sub>2</sub>	ppb	0.0143	0.2540
		(NO <sub>2</sub> ) <sup>2</sup>	ppb <sup>2</sup>	-0.0003	0.5968
		(NO <sub>2</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.6912
		SO <sub>2</sub>	ppb	-0.0030	0.6781
		(SO <sub>2</sub> ) <sup>2</sup>	ppb <sup>2</sup>	0.0002	0.6257
		(SO <sub>2</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.6899
Secondary	Model including pollutants in joint effect, also controlling for CO, NO <sub>2</sub> , and SO <sub>2</sub> , no interactions	O <sub>3</sub>	ppb	0.0006	0.5886
		CO	ppm	0.0895	0.0754
		NO <sub>2</sub>	ppb	0.0039	0.0859
		SO <sub>2</sub>	ppb	0.0001	0.9460
		Secondary PM <sub>2.5</sub> <sup>b</sup>	µg/m <sup>3</sup>	0.0005	0.8704
	Model including only pollutants in joint effect, with linear, quadratic and cubic terms for each pollutant	O <sub>3</sub>	ppb	-0.0074	0.3461
		(O <sub>3</sub> ) <sup>2</sup>	ppb <sup>2</sup>	0.0002	0.2095
		(O <sub>3</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.2360
		Secondary PM <sub>2.5</sub> <sup>b</sup>	µg/m <sup>3</sup>	0.0033	0.8755
		(Secondary PM <sub>2.5</sub> ) <sup>2b</sup>	(µg/m <sup>3</sup> ) <sup>2</sup>	-0.0004	0.8110
(Secondary PM <sub>2.5</sub> ) <sup>3b</sup>	(µg/m <sup>3</sup> ) <sup>3</sup>	0.0000	0.7255		
Traffic	Model including pollutants in joint effect, also controlling for O <sub>3</sub> , SO <sub>2</sub> and SO <sub>4</sub> <sup>2-</sup> , no interactions	O <sub>3</sub>	ppb	0.0008	0.4801
		CO	ppm	0.1179	0.0430
		NO <sub>2</sub>	ppb	0.0041	0.0706
		SO <sub>2</sub>	ppb	0.0001	0.9468
		EC	µg/m <sup>3</sup>	-0.0369	0.3133
		SO <sub>4</sub> <sup>2-</sup>	µg/m <sup>3</sup>	0.0006	0.8823
	Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	CO	ppm	0.7011	0.0347
		(CO) <sup>2</sup>	ppm <sup>2</sup>	-0.5832	0.0520
		(CO) <sup>3</sup>	ppm <sup>3</sup>	0.1580	0.0466
		NO <sub>2</sub>	ppb	0.0155	0.2502
		(NO <sub>2</sub> ) <sup>2</sup>	ppb <sup>2</sup>	-0.0005	0.3825
		(NO <sub>2</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.3589
		EC	µg/m <sup>3</sup>	-0.2653	0.0611
		(EC) <sup>2</sup>	(µg/m <sup>3</sup> ) <sup>2</sup>	0.2277	0.0243
(EC) <sup>3</sup>	(µg/m <sup>3</sup> ) <sup>3</sup>	-0.0572	0.0057		
Power Plant	Model including pollutants in joint effect , also controlling for O <sub>3</sub> , CO, NO <sub>2</sub> and EC, no interactions	O <sub>3</sub>	ppb	0.0008	0.4801
		CO	ppm	0.1179	0.0430
		NO <sub>2</sub>	ppb	0.0041	0.0706
		SO <sub>2</sub>	ppb	0.0001	0.9468
		EC	µg/m <sup>3</sup>	-0.0369	0.3133
		SO <sub>4</sub> <sup>2-</sup>	µg/m <sup>3</sup>	0.0006	0.8823
	Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	SO <sub>2</sub>	ppb	0.0104	0.1276
		(SO <sub>2</sub> ) <sup>2</sup>	ppb <sup>2</sup>	-0.0004	0.3559
		(SO <sub>2</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.4906
		SO <sub>4</sub> <sup>2-</sup>	µg/m <sup>3</sup>	0.0101	0.6992
(SO <sub>4</sub> <sup>2-</sup> ) <sup>2</sup>	(µg/m <sup>3</sup> ) <sup>2</sup>	-0.0008	0.8132		
(SO <sub>4</sub> <sup>2-</sup> ) <sup>3</sup>	(µg/m <sup>3</sup> ) <sup>3</sup>	0.0000	0.7686		
Criteria Pollutants	Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	O <sub>3</sub>	ppb	-0.0072	0.3505
		(O <sub>3</sub> ) <sup>2</sup>	ppb <sup>2</sup>	0.0002	0.3141
		(O <sub>3</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.3600
		CO	ppm	0.5136	0.1138
		(CO) <sup>2</sup>	ppm <sup>2</sup>	-0.3857	0.1950
		(CO) <sup>3</sup>	ppm <sup>3</sup>	0.1036	0.2096
		NO <sub>2</sub>	ppb	0.0081	0.5549
		(NO <sub>2</sub> ) <sup>2</sup>	ppb <sup>2</sup>	-0.0002	0.7217
		(NO <sub>2</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.7645
		SO <sub>2</sub>	ppb	-0.0009	0.9020
		(SO <sub>2</sub> ) <sup>2</sup>	ppb <sup>2</sup>	0.0001	0.7966
		(SO <sub>2</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.7096
		PM <sub>2.5</sub>	µg/m <sup>3</sup>	-0.0089	0.6576
		(PM <sub>2.5</sub> ) <sup>2</sup>	(µg/m <sup>3</sup> ) <sup>2</sup>	0.0003	0.7950
		(PM <sub>2.5</sub> ) <sup>3</sup>	(µg/m <sup>3</sup> ) <sup>3</sup>	0.0000	0.8969

Abbreviations: O<sub>3</sub>=ozone, CO=carbon monoxide, NO<sub>2</sub>=nitrogen dioxide, SO<sub>2</sub>=sulfur dioxide, PM<sub>2.5</sub>=particulate matter < 2.5 µm in diameter, EC=elemental carbon component of PM<sub>2.5</sub>, SO<sub>4</sub><sup>2-</sup>=sulfate component of PM<sub>2.5</sub>, ppb=parts per billion, ppm=parts per million, µg/m<sup>3</sup>=micrograms per cubic millimeter.

<sup>a</sup> Shaded pollutants were not included in calculation of joint effect.

<sup>b</sup> Secondary PM<sub>2.5</sub> was calculated as the sum of the concentrations of the PM<sub>2.5</sub> components sulfate, nitrate, and ammonium.

**eTable 10. Pollutant-specific parameter estimates from sensitivity analyses, Atlanta, 1998-2004, Cold Season**

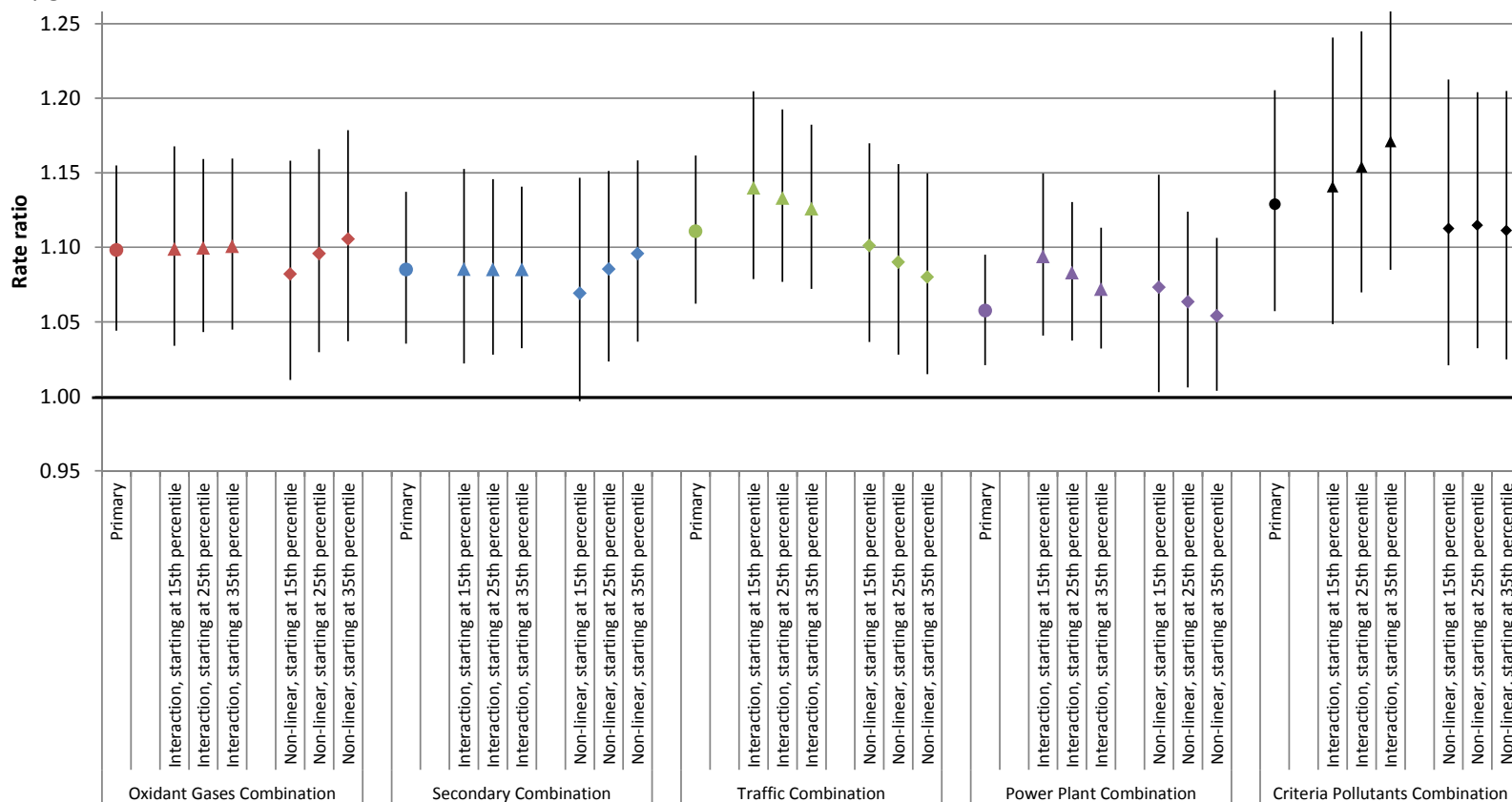
Joint Effect	Model Specifications	Pollutant <sup>a</sup>	Unit	Coefficient per unit change	p-value
Oxidant Gases	Model including pollutants in joint effect, also controlling for CO and PM <sub>2.5</sub> , no interactions	O <sub>3</sub>	ppb	0.0017	0.1539
		CO	ppm	0.0078	0.8306
		NO <sub>2</sub>	ppb	0.0022	0.2942
		SO <sub>2</sub>	ppb	-0.0020	0.1536
		PM <sub>2.5</sub>	µg/m <sup>3</sup>	-0.0024	0.2096
	Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	O <sub>3</sub>	ppb	-0.0160	0.0491
		(O <sub>3</sub> ) <sup>2</sup>	ppb <sup>2</sup>	0.0004	0.0579
		(O <sub>3</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.1302
		NO <sub>2</sub>	ppb	-0.0046	0.7993
		(NO <sub>2</sub> ) <sup>2</sup>	ppb <sup>2</sup>	0.0002	0.7776
		(NO <sub>2</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.7914
		SO <sub>2</sub>	ppb	0.0060	0.4683
		(SO <sub>2</sub> ) <sup>2</sup>	ppb <sup>2</sup>	-0.0005	0.3280
		(SO <sub>2</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.3088
Secondary	Model including pollutants in joint effect, also controlling for CO, NO <sub>2</sub> , and SO <sub>2</sub> , no interactions	O <sub>3</sub>	ppb	0.0021	0.0789
		CO	ppm	-0.0025	0.9464
		NO <sub>2</sub>	ppb	0.0023	0.2570
		SO <sub>2</sub>	ppb	-0.0018	0.2255
		Secondary PM <sub>2.5</sub> <sup>b</sup>	µg/m <sup>3</sup>	-0.0061	0.1194
	Model including only pollutants in joint effect, with linear, quadratic and cubic terms for each pollutant	O <sub>3</sub>	ppb	-0.0183	0.0228
		(O <sub>3</sub> ) <sup>2</sup>	ppb <sup>2</sup>	0.0005	0.0226
		(O <sub>3</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.0656
		Secondary PM <sub>2.5</sub> <sup>b</sup>	µg/m <sup>3</sup>	-0.0306	0.2433
		(Secondary PM <sub>2.5</sub> ) <sup>2b</sup>	(µg/m <sup>3</sup> ) <sup>2</sup>	0.0047	0.1341
(Secondary PM <sub>2.5</sub> ) <sup>3b</sup>	(µg/m <sup>3</sup> ) <sup>3</sup>	-0.0002	0.0410		
Traffic	Model including pollutants in joint effect, also controlling for O <sub>3</sub> , SO <sub>2</sub> and SO <sub>4</sub> <sup>2-</sup> , no interactions	O <sub>3</sub>	ppb	0.0022	0.0633
		CO	ppm	-0.0062	0.8736
		NO <sub>2</sub>	ppb	0.0021	0.3260
		SO <sub>2</sub>	ppb	-0.0017	0.2607
		EC	µg/m <sup>3</sup>	0.0021	0.9319
		SO <sub>4</sub> <sup>2-</sup>	µg/m <sup>3</sup>	-0.0072	0.2392
	Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	CO	ppm	0.2945	0.2740
		(CO) <sup>2</sup>	ppm <sup>2</sup>	-0.2698	0.2277
		(CO) <sup>3</sup>	ppm <sup>3</sup>	0.0703	0.2183
		NO <sub>2</sub>	ppb	-0.0104	0.5815
		(NO <sub>2</sub> ) <sup>2</sup>	ppb <sup>2</sup>	0.0004	0.5013
		(NO <sub>2</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.5332
		EC	µg/m <sup>3</sup>	-0.0730	0.4805
		(EC) <sup>2</sup>	(µg/m <sup>3</sup> ) <sup>2</sup>	0.0327	0.5735
(EC) <sup>3</sup>	(µg/m <sup>3</sup> ) <sup>3</sup>	-0.0046	0.6078		
Power Plant	Model including pollutants in joint effect , also controlling for O <sub>3</sub> , CO, NO <sub>2</sub> and EC, no interactions	O <sub>3</sub>	ppb	0.0022	0.0633
		CO	ppm	-0.0062	0.8736
		NO <sub>2</sub>	ppb	0.0021	0.3260
		SO <sub>2</sub>	ppb	-0.0017	0.2607
		EC	µg/m <sup>3</sup>	0.0021	0.9319
		SO <sub>4</sub> <sup>2-</sup>	µg/m <sup>3</sup>	-0.0072	0.2392
	Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	SO <sub>2</sub>	ppb	0.0100	0.2223
		(SO <sub>2</sub> ) <sup>2</sup>	ppb <sup>2</sup>	-0.0008	0.1487
		(SO <sub>2</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.1641
		SO <sub>4</sub> <sup>2-</sup>	µg/m <sup>3</sup>	-0.0404	0.2318
		(SO <sub>4</sub> <sup>2-</sup> ) <sup>2</sup>	(µg/m <sup>3</sup> ) <sup>2</sup>	0.0109	0.0846
(SO <sub>4</sub> <sup>2-</sup> ) <sup>3</sup>	(µg/m <sup>3</sup> ) <sup>3</sup>	-0.0007	0.0225		
Criteria Pollutants	Model including only pollutants in joint effects, with linear, quadratic and cubic terms for each pollutant	O <sub>3</sub>	ppb	-0.0172	0.0362
		(O <sub>3</sub> ) <sup>2</sup>	ppb <sup>2</sup>	0.0005	0.0428
		(O <sub>3</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.0917
		CO	ppm	0.3685	0.1537
		(CO) <sup>2</sup>	ppm <sup>2</sup>	-0.3269	0.1305
		(CO) <sup>3</sup>	ppm <sup>3</sup>	0.0838	0.1326
		NO <sub>2</sub>	ppb	-0.0132	0.4931
		(NO <sub>2</sub> ) <sup>2</sup>	ppb <sup>2</sup>	0.0005	0.4662
		(NO <sub>2</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.5101
		SO <sub>2</sub>	ppb	0.0114	0.1800
		(SO <sub>2</sub> ) <sup>2</sup>	ppb <sup>2</sup>	-0.0010	0.0935
		(SO <sub>2</sub> ) <sup>3</sup>	ppb <sup>3</sup>	0.0000	0.0847
		PM <sub>2.5</sub>	µg/m <sup>3</sup>	-0.0253	0.0625
		(PM <sub>2.5</sub> ) <sup>2</sup>	(µg/m <sup>3</sup> ) <sup>2</sup>	0.0013	0.0374
		(PM <sub>2.5</sub> ) <sup>3</sup>	(µg/m <sup>3</sup> ) <sup>3</sup>	0.0000	0.0168

Abbreviations: O<sub>3</sub>=ozone, CO=carbon monoxide, NO<sub>2</sub>=nitrogen dioxide, SO<sub>2</sub>=sulfur dioxide, PM<sub>2.5</sub>=particulate matter < 2.5 µm in diameter, EC=elemental carbon component of PM<sub>2.5</sub>, SO<sub>4</sub><sup>2-</sup>=sulfate component of PM<sub>2.5</sub>, ppb=parts per billion, ppm=parts per million, µg/m<sup>3</sup>=micrograms per cubic millimeter.

<sup>a</sup> Shaded pollutants were not included in calculation of joint effect.

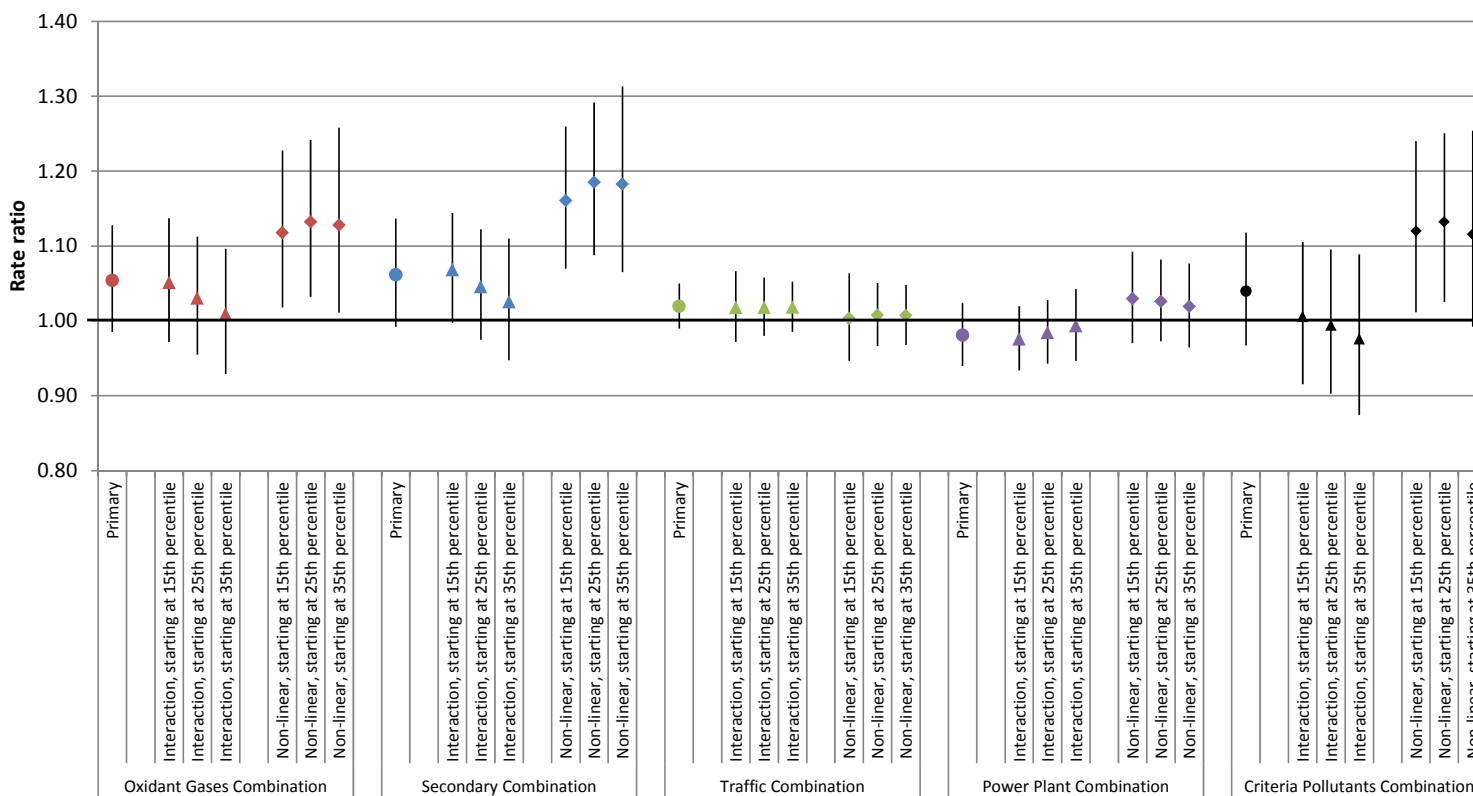
<sup>b</sup> Secondary PM<sub>2.5</sub> was calculated as the sum of the concentrations of the PM<sub>2.5</sub> components sulfate, nitrate, and ammonium.

**eFigure 1. Rate ratios for joint effects from multipollutant models, evaluated for various specific IQR increments in each pollutant, Warm Season.** Rate ratios are shown in red for the oxidant pollutant combination, blue for the secondary pollutant combination, green for the traffic pollutant combination, purple for the power plant pollutant combination, and black for the criteria pollutant combination. For each combination of pollutants, joint effect rate ratios from the primary multipollutant models, which had linear pollutant terms and no interactions (“Primary”, circle markers), are followed by multipollutant models with linear pollutant terms and first order interactions between pollutants, with IQR changes evaluated starting at the 15<sup>th</sup>, 25<sup>th</sup>, and 35<sup>th</sup> percentiles for each pollutant (“Interaction”, triangle markers); and multipollutant models with linear, quadratic and cubic pollutant terms and no interactions, with IQR changes evaluated starting at the 15<sup>th</sup>, 25<sup>th</sup>, and 35<sup>th</sup> percentiles for each pollutant (“Non-linear”, diamond markers). For comparability with previous analyses,<sup>1</sup> the analysis used the year-round IQR during 1993-2004 for O<sub>3</sub>, NO<sub>2</sub>, CO and SO<sub>2</sub>, and during August 1, 1998-December 31, 2004 for PM<sub>2.5</sub> and PM<sub>2.5</sub> components. The IQRs used in the analysis were: O<sub>3</sub> 29.18 ppb, CO 0.66 ppm, NO<sub>2</sub> 12.87 ppb, SO<sub>2</sub> 10.51 ppb, PM<sub>2.5</sub> 9.18 µg/m<sup>3</sup>, EC 0.69 µg/m<sup>3</sup>, SO<sub>4</sub><sup>2-</sup> 3.45 µg/m<sup>3</sup>, Secondary PM<sub>2.5</sub> 4.52 µg/m<sup>3</sup>.

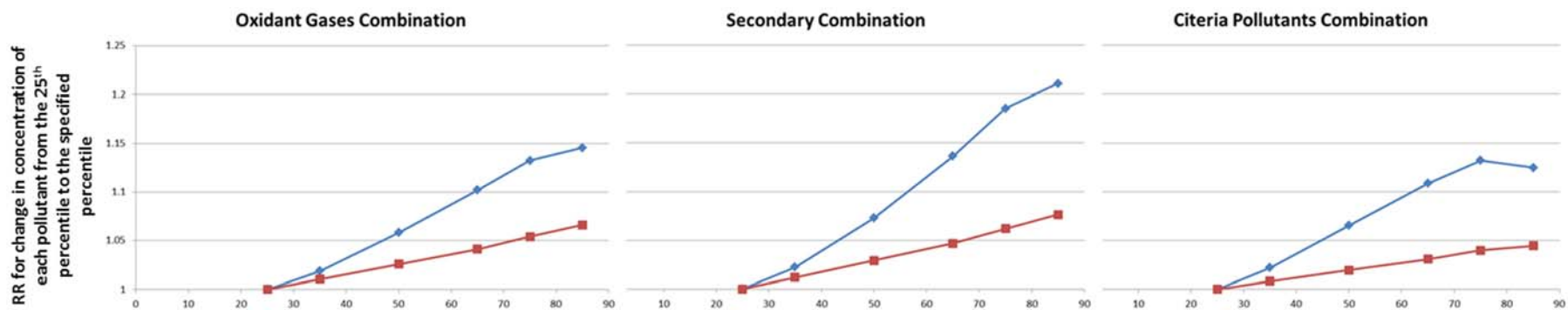




**eFigure 2. Rate ratios for joint effects from multipollutant models, evaluated for various specific IQR increments in each pollutant, Cold Season.** Rate ratios are shown in red for the oxidant pollutant combination, blue for the secondary pollutant combination, green for the traffic pollutant combination, purple for the power plant pollutant combination, and black for the criteria pollutant combination. For each combination of pollutants, joint effect rate ratios from the primary multipollutant models, which had linear pollutant terms and no interactions (“Primary”, circle markers), are followed by multipollutant models with linear pollutant terms and first order interactions between pollutants, with IQR changes evaluated starting at the 15<sup>th</sup>, 25<sup>th</sup>, and 35<sup>th</sup> percentiles for each pollutant (“Interaction”, triangle markers); and multipollutant models with linear, quadratic and cubic pollutant terms and no interactions, with IQR changes evaluated starting at the 15<sup>th</sup>, 25<sup>th</sup>, and 35<sup>th</sup> percentiles for each pollutant (“Non-linear”, diamond markers). For comparability with previous analyses,<sup>1</sup> the analysis used the year-round IQR during 1993-2004 for O<sub>3</sub>, NO<sub>2</sub>, CO and SO<sub>2</sub>, and during August 1, 1998-December 31, 2004 for PM<sub>2.5</sub> and PM<sub>2.5</sub> components. The IQRs used in the analysis were: O<sub>3</sub> 29.18 ppb, CO 0.66 ppm, NO<sub>2</sub> 12.87 ppb, SO<sub>2</sub> 10.51 ppb, PM<sub>2.5</sub> 9.18 µg/m<sup>3</sup>, EC 0.69 µg/m<sup>3</sup>, SO<sub>4</sub><sup>2-</sup> 3.45 µg/m<sup>3</sup>, Secondary PM<sub>2.5</sub> 4.52 µg/m<sup>3</sup>.



eFigure 3. Rate ratios per changes in the concentration of each pollutant from the 25<sup>th</sup> percentile to various specified percentiles based on results of joint effect models with and without non-linear terms (quadratic and cubic pollutant terms), for selected pollutant combinations, Atlanta, 1998-2004, cold season.



Percentile of each pollutant distribution (absolute concentrations for the percentiles for each pollutant are listed below)

		With Non-linear terms						Only linear terms											
Percentile		25	35	50	65	75	85	25	35	50	65	75	85	25	35	50	65	75	85
Absolute pollutant concentrations*	O3 (ppb)	30.10	35.44	43.55	52.02	59.27	67.44	30.10	35.44	43.55	52.02	59.27	67.44	30.10	35.44	43.55	52.02	59.27	67.44
	CO (ppb)													0.54	0.63	0.78	0.99	1.19	1.45
	NO2 (ppb)	16.29	18.68	22.04	26.04	29.16	32.89	16.29	18.68	22.04	26.04	29.16	32.89	16.29	18.68	22.04	26.04	29.16	32.89
	SO2 (ppb)	4.03	5.49	8.00	11.45	14.54	19.48	4.03	5.49	8.00	11.45	14.54	19.48	4.03	5.49	8.00	11.45	14.54	19.48
	PM2.5 (µg/m <sup>3</sup> )													10.96	12.53	14.97	17.71	20.14	24.01
	Secondary PM (µg/m <sup>3</sup> )							4.26	4.91	6.05	7.48	8.77	10.82						

\*For comparability with previous analyses,<sup>1</sup> the percentiles were based on the year-round distribution during 1993-2004 for O3, NO2, CO and SO2, and during August 1, 1998-December 31, 2004 for PM2.5 and PM2.5 components.

## References

1. Strickland MJ, Darrow LA, Klein M, et al. Short-term Associations between Ambient Air Pollutants and Pediatric Asthma Emergency Department Visits. *Am J Respir Crit Care Med* 2010;182:307-16.